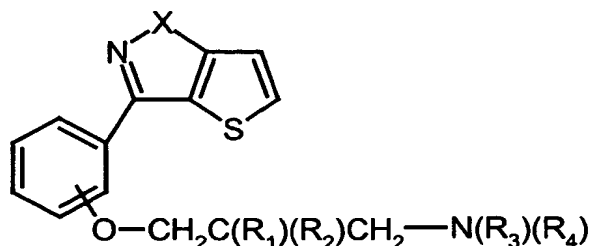


What is claimed is:

1. A compound of Formula I :



Formula I

- 5 a pharmaceutically acceptable salt or stereoisomer thereof,

wherein

X is N(CH₃) or O;

R₁ is OH or C₁₋₆ alkoxy;

R₂ is H or C₁₋₆ alkyl;

- 10 R₃ is (CH₂)_n Q, CH₂CH(OH)Q, CH(CH₃)Q, 1,2,3,4-tetrahydronaphthyl, indanyl, or adamantyl, wherein

Q is thienyl, phenyl, furanyl, naphthyl, pyridyl, indolyl, indazolyl, cyclohexyl, 1,2-methylenedioxyphenyl, cyclohexenyl, 1H-pyrazolo[4,3-c]pyridyl, and

- 15 Q is optionally substituted with one or two moieties independently selected from halo, C₁₋₆ alkyl, C₁₋₆ alkoxy, hydroxy, S(O)₂NH₂, trifluoromethyl, or cyano, and

n is 1 or 2;

- 20 R₄ is H or C₁₋₆ alkyl; or

R₃ and R₄, together with the nitrogen atom to which R₃ and R₄ are attached, form 1,4-dioxo-8-azo-spiro[4.5]decanyl, piperazinyl, morpholinyl, piperidinyl, pyrrolidinyl, azocanyl, 1,2,3,4-tetrahydroisoquinolinyl, 1,2,3,4-tetrahydro-β-carbolinyl, 4,5,6,7-tetrahydrothienyl[3,2-c]pyridyl, or 8-aza-bicyclo[3.2.1]octane, each of which may be mono- or independently di-substituted with halo, C₁₋₆ alkyl, C₁₋₆ alkoxy, C(O)phenyl, OH, CN, O-phenyl or (CH₂)_mZ,

25

Z is benzisoxazolyl, indazolyl, benzisothiazolyl, benzothienyl, pyrimidinyl, pyridyl, 1,2-methylenedioxyphenyl, or phenyl, and

Z, CH(OH)phenyl or O-phenyl are optionally substituted with one or two moieties independently selected from halo, C₁₋₆ alkyl, C₁₋₆ alkoxy, hydroxy, trifluoromethyl, S(O)₂NH₂, or cyano, and

m is 0 or 1;

provided that when R₁ is OH, R₂ is H:

- (1) R₄ is H, and R₃ is (CH₂)_nQ, where n is 1 or 2, then Q cannot be indolyl or phenyl; or
 - (2) R₃ and R₄ form piperazinyl substituted with (CH₂)_mZ, when m is 1, then Z cannot be phenyl.
2. A compound according to claim 1 wherein Q is thienyl or pyridyl; or R₃ and R₄, together with the nitrogen atom to which R₃ and R₄ are attached, form piperidinyl.
 3. The compound according to claim 2 which is (2R)-2-methyl-1-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-3-[thienophen-2-ylmethyl]-amino-propan-2-ol.
 4. The compound according to claim 2 which is (2R)-2-methyl-1-[(pyridin-3-ylmethyl)-amino]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
 5. The compound according to claim 2 which is (2R)-2-methyl-1-[(pyridin-2-ylmethyl)-amino]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
 6. The compound according to claim 2 which is (2S)-1-(2-thienylmethylamino)-2-methyl-3-[3-thieno[2,3-d]isoxazol-3-yl-phenoxy]propan-2-ol.
 7. The compound according to claim 2 which is (2S)-1-[4-(3-chlorophenoxy)-1-piperidinyl]-2-methyl-3-[3-thieno[2,3-d]isoxazol-3-yl-phenoxy]propan-2-ol.

8. The compound according to claim 2 which is (2S)-1-[4-(6-fluorobenzo[d]isoxazol-3-yl)piperidin-1-yl]-2-methyl-3-[3-thieno[2,3-d]isoxazol-3-yl-phenoxy]propan-2-ol.
9. The compound according to claim 2 which is (2R)-2-methyl-1-[(pyridin-4-ylmethyl)-amino]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
10. The compound according to claim 2 which is (2R)-1-[4-(6-fluorobenzo[d]isoxazol-3-yl)piperidin-1-yl]-3-(4-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
11. The compound according to claim 2 which is (2S)-1-(3-thienylmethylamino)-2-methyl-3-[3-thieno[2,3-d]isoxazol-3-yl-phenoxy]propan-2-ol.
12. The compound according to claim 2 which is (2S)-2-methyl-1-[(pyridin-3-ylmethyl)amino]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
13. The compound according to claim 2 which is (2S)-2-methyl-1-[(pyridin-2-ylmethyl)amino]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
14. The compound according to claim 2 which is (4-fluorophenyl)-(1-{2(R)-hydroxy-3-[3-(1-methyl-1H-thieno[3,2-c]pyrazol-3-yl)-phenoxy]-propyl}-piperidin-4-yl)-methanone.
15. The compound according to claim 2 which is 1-(1-{2(R)-hydroxy-3-[3-(1-methyl-1H-thieno[3,2-c]pyrazol-3-yl)-phenoxy]-propyl}-piperidin-4-yl)-1,3-dihydrobenzimidazol-2-one.
16. A compound according to claim 2 wherein
 R_1 is OH;
 R_2 is H;
 R_3 is $(CH_2)_n$ Q; or
 R_3 and R_4 together with the nitrogen atom to which R_3 and R_4 are attached form piperidiny1; and
 n is 1.
17. A compound according to claim 16 wherein Q is thienyl.
18. The compound according to claim 17 which is (2R)-1-(4-thieno[2,3-d]isoxazol-3-yl-phenoxy)-3-[(thiophen-3-ylmethyl)-amino]-propan-2-ol.
19. The compound of claim 17 which is (2R)-1-(4-thieno[2,3-d]isoxazol-3-yl-phenoxy)-3-[(thiophen-2-ylmethyl)-amino]-propan-2-ol.

20. The compound of claim 17 which is (2R)-1-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-3-[(thiophen-2-ylmethyl)amino]propan-2-ol.
21. The compound of claim 17 which is (2R)-1-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-3-[(thiophen-3-ylmethyl)amino]propan-2-ol.
- 5 22. A compound according to claim 16 wherein Q is pyridyl.
23. The compound of claim 22 which is (2R)-1-[(pyridin-4-yl)methylamino]-3-[3-thieno[2,3-d]isoxazol-3-yl-phenoxy]propan-2-ol.
24. The compound of claim 22 which is (2R)-1-[(pyridin-2-ylmethyl)-amino]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
- 10 25. The compound of claim 22 which is (2R)-1-[(pyridin-3-ylmethyl)amino]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
26. A compound according to claim 16 wherein R₃ and R₄, together with the nitrogen atom to which R₃ and R₄ are attached, form piperidiny1.
27. The compound according to claim 26 which is 4-(4-chlorophenyl)-1-[(R)-2-hydroxy-3-[3-(thieno[2,3-d]isoxazol-3-yl)phenoxy]propyl]piperidin-4-ol.
- 15 28. The compound of claim 26 which is (2R)-1-[4-(6-fluorobenzo[d]isoxazol-3-yl)piperidin-1-yl]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
29. The compound of claim 26 which is (2R)-1-[4-(6-chlorobenzo[d]isoxazol-3-yl)piperidin-1-yl]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
- 20 30. The compound of claim 26 which is (2R)-1-[4-(6-fluorobenzo[d]isothiazol-3-yl)piperidin-1-yl]-3-(thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
31. The compound of claim 26 which is (2R)-1-(4-benzylpiperidin-1-yl)-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
32. The compound of claim 26 which is (2R)-1-piperidin-1-yl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
- 25 33. A compound according to claim 1 wherein
R₃ and R₄, together with the nitrogen atom to which R₃ and R₄ are attached, form piperaziny1.
34. The compound of claim 33 which is (2R)-1-[4-(4-chlorophenyl)piperazin-1-yl]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
- 30 35. The compound of claim 33 which is (2R)-1-[4-(2-methoxyphenyl)-piperazin-1-yl]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.

36. The compound of claim 33 which is (2R)-1-[4-(2-fluorophenyl)piperazin-1-yl]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
37. The compound of claim 33 which is (2R)-1-[4-(4-fluorophenyl)piperzain-1-yl]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
- 5 38. The compound of claim 33 which is (2R)-1-[4-(2-chlorophenyl)piperazin-1-yl]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
39. The compound of claim 33 which is (2R)-1-[4-(3-chlorophenyl)piperazin-1-yl]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
40. The compound of claim 33 which is (2R)-1-[4-(4-methoxyphenyl)piperazin-1-yl]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
- 10 41. The compound of claim 33 which is (2R)-1-(4-phenylpiperazin-1-yl)-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
42. The compound of claim 33 which is 2-{4-[(R)-2-hydroxy-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propyl]piperazin-1-yl}benzonitrile.
- 15 43. The compound of claim 33 which is (2R)-1-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-3-[4-(2-trifluoromethylphenyl)-piperazin-1-yl]propan-2-ol.
44. The compound of claim 33 which is (2R)-1-[4-(2-methoxyphenyl)piperazin-1-yl]-3-(4-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
45. The compound of claim 33 which is 2-{4-[(R)-2-hydroxy-3-(4-thieno[2,3-d]isoxazol-3-yl-phenoxy)propyl]-piperazin-1-yl}benzonitrile.
- 20 46. The compound of claim 33 which is (2S)-1-[4-(2-methoxyphenyl)piperazin-1-yl]-2-methyl-3-[3-thieno[2,3-d]isoxazol-3-yl-phenoxy]propan-2-ol.
47. The compound of claim 33 which is (2S)-1-[4-(2-cyanophenyl)-1-piperazinyl]-2-methyl-3-[3-thieno[2,3-d]isoxazol-3-yl-phenoxy]propan-2-ol.
- 25 48. The compound of claim 33 which is (2R)-1-(4-pyrimidin-2-yl-piperazin-1-yl)-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
49. The compound of claim 33 which is (2R)-1-(4-pyridin-2-yl-piperazin-1-yl)-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
50. The compound of claim 33 which is (2R)-1-(4-benzo[1,3]dioxol-5-ylmethyl-piperazin-1-yl)-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
- 30 51. The compound of claim 33 which is (2R)-1-[4-(6-fluoro-1H-indazol-3-yl)-piperazin-1-yl]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.

52. The compound of claim 33 which is (2R)-1-[4-(5-methoxy-1H-indazol-3-yl)-piperazin-1-yl]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
53. The compound of claim 33 which is (2R)-1-(4-benzo[d]isothiazol-3-yl-piperazin-1-yl)-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
54. The compound of claim 33 which is (2R)-1-[4-(6-fluorobenzo[b]thiophen-3-yl)piperazin-1-yl]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
55. The compound of claim 33 which is 3-[(R)-4-[2-hydroxy-3-(3-thieno[2,3-d]isoxazol-3-yl)phenoxy]propyl]-piperazin-1-yl]-benzo[d]isoxazol-6-ol.
56. The compound of claim 33 which is (2R)-1-[4-(4-methoxyphenyl)-3-methylpiperazin-1-yl]-3-[3-(thieno[2,3-d]isoxazol-3-yl)phenoxy]propan-2-ol.
57. The compound of claim 33 which is (2R)-1-[3-(1-methyl-1H-thieno[3,2-c]pyrazol-3-yl)-phenoxy-3-(4-phenyl-piperazin-1-yl)-propan-2-ol.
58. The compound of claim 33 which is (2R)-1-[3-(1-methyl-1H-thieno[3,2-c]pyrazol-3-yl)-phenoxy-3-(4-pyrimidin-2-yl-piperazin-1-yl)-propan-2-ol.
59. A compound according to claim 1 wherein Q is phenyl.
60. The compound of claim 59 which is (2R)-1-(4-chlorobenzylamino)-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
61. The compound of claim 59 which is (2R)-1-[(N-benzyl-N-methyl)amino]-3-[2-thieno[2,3-d]isoxazol-3-yl)phenoxy]-2-propanol.
62. The compound of claim 59 which is (2S)-(+)-1-benzylamino-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
63. The compound of claim 59 which is (2R)-(-)-1-benzylamino-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
64. The compound of claim 59 which is (2R)-1-(benzylmethylamino)-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
65. The compound of claim 59 which is (2R)-1-(4-methoxybenzylamino)-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
66. The compound of claim 59 which is (2R)-1-(4-fluorobenzylamino)-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
67. The compound of claim 59 which is (2R)-1-(2-fluorobenzylamino)-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
68. The compound of claim 59 which is (2R)-2-methyl-1-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-3-(4-trifluoromethylbenzylamino)-propan-2-ol.

69. The compound of claim 59 which is (2R)-2-methyl-1-[1(R)-phenylethylamino]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
70. The compound of claim 59 which is (2R)-2-methyl-1-[1(S)-phenylethylamino]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
- 5 71. The compound of claim 59 which is (2R)-1-(2-hydroxy-2-phenylethylamino)-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
72. The compound of claim 59 which is (2R)-benzyl-[2-methoxy-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propyl]-methylamine hydrochloride.
73. The compound of claim 59 which is (2R)-benzyl-[2-methoxy-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propyl]-amine.
- 10 74. The compound of claim 59 which is (2S)-1-(4-fluorobenzylamino)-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
75. The compound of claim 59 which is (2S)-1-(2-fluorobenzylamino)-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
- 15 76. The compound of claim 59 which is (2S)-1-(3-fluorobenzylamino)-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
77. The compound of claim 59 which is (2S)-1-(4-chlorobenzylamino)-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
78. The compound of claim 59 which is (2S)-1-(2-chlorobenzylamino)-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
- 20 79. The compound of claim 59 which is (2S)-1-(3,4-dichlorobenzylamino)-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
80. The compound of claim 59 which is (2S)-2-methyl-1-[1(R)-phenylethylamino]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
- 25 81. The compound of claim 59 which is (2S)-2-methyl-1-[1(S)-phenylethylamino]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
82. The compound of claim 59 which is (2S)-2-methyl-1-(4-methylbenzylamino)-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
83. The compound of claim 59 which is (2S)-1-(4-methoxybenzylamino)-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
- 30 84. The compound of claim 59 which is (2S)-1-(2-methoxybenzylamino)-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.

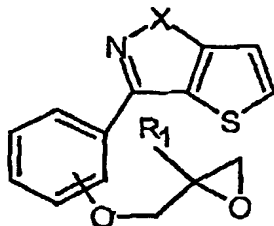
85. The compound of claim 59 which is (2S)-1-(benzylmethylamino)-2-methyl-3-[3-thieno[2,3-d]isoxazol-3-yl-phenoxy]propan-2-ol.
86. The compound of claim 59 which is (2S)-1-(3,4-difluorobenzylamino)-2-methyl-3-[3-(thieno[2,3-d]isoxazol-3-yl)phenoxy]propan-2-ol.
- 5 87. The compound of claim 59 which is (2R)-1-(2-methoxybenzylamino)-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
88. A compound according to claim 1 wherein Q is furanyl.
89. The compound of claim 88 which is (2R)-1-[(furan-2-ylmethyl)-amino]-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
- 10 90. The compound of claim 88 which is (2R)-1-[(furan-2-ylmethyl)amino]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
91. The compound of claim 88 which is (2R)-1-[(furan-2-ylmethyl)-amino]-3-(4-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
92. The compound of claim 88 which is (2S)-1-[(furan-2-ylmethyl)amino]-2-methyl-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
- 15 93. A compound according to claim 1 wherein R₃ is indanyl.
94. The compound of claim 93 which is (2R)-1-(indan-1-ylamino)-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
95. The compound of claim 93 which is (2R)-1-(indan-2-ylamino)-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
- 20 96. A compound according to claim 1 wherein Q is naphthyl.
97. The compound of claim 96 which is (2R)-1-[(naphthalen-1-ylmethyl)amino]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
98. The compound of claim 96 which is (2R)-2-methyl-1-[(naphthalen-1-ylmethyl)amino]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
- 25 99. The compound of claim 96 which is (2R)-1-[(naphthalen-1-ylmethyl)-amino]-3-(4-thieno[2,3-d]isoxazol-3-yl-phenoxy)-propan-2-ol.
100. The compound of claim 96 which is (2S)-2-methyl-1-[(naphthalen-1-ylmethyl)amino]-3-(3-thieno[2,3-d]isoxazol-3-yl-phenoxy)propan-2-ol.
- 30 101. A compound according to claim 1 wherein R₃ and R₄, together with the nitrogen atom to which R₃ and R₄ are attached, form 1,2,3,4-tetrahydroisoquinoliny.

102. The compound of claim 101 which is (\pm) -1-(3,4-dihydro-1H-isoquinolin-2-yl)-3-[2-(thieno[2,3-*d*]isoxazol-3-yl)phenoxy]-2-propanol.
103. The compound of claim 101 which is (2R)-1-(6,7-dimethoxy-1,2,3,4-tetrahydroisoquinolin-2-yl)-3-[3-(thieno[2,3-*d*]isoxazol-3-yl)phenoxy]propan-2-ol.
104. The compound of claim 101 which is (2R)-1-(3,4-dihydro-1H-isoquinolin-2-yl)-3-[2-(thieno[2,3-*d*]isoxazol-3-yl)phenoxy]-2-propanol.
105. The compound of claim 101, which is (2R)-1-(3,4-dihydro-1H-isoquinolin-2-yl)-3-[3-(1-methyl-1H-thieno[3,2-*c*]pyrazol-3-yl)-phenoxy]-propan-2-ol.
106. A compound according to claim 1, wherein R_3 and R_4 , together with the nitrogen atom to which R_3 and R_4 are attached, form 1,2,3,4-tetrahydro- β -carbolinyl.
107. The compound of claim 106 which is (2R)-1-(1,2,3,4-tetrahydro- β -carbolin-2-yl)-3-[3-thieno[2,3-*d*]isoxazol-3-yl)phenoxy]propan-2-ol.
108. A compound according to claim 1, wherein R_3 and R_4 , together with the nitrogen atom to which R_3 and R_4 are attached, form 4,5,6,7-tetrahydrothieno[3,2-*c*]pyridinyl.
109. The compound of claim 108 which is (2R)-1-(6,7-dihydro-4H-thieno[3,2-*c*]pyridin-5-yl)-3-(3-thieno[2,3-*d*]isoxazol-3-yl-phenoxy)propan-2-ol.
110. The compound of claim 108 which is (2R)-1-(6,7-dihydro-4H-thieno[3,2-*c*]pyridin-5-yl)-3-(4-thieno[2,3-*d*]isoxazol-3-yl-phenoxy)propan-2-ol.
111. A compound according to claim 1, wherein R_3 and R_4 , together with the nitrogen atom to which R_3 and R_4 are attached, form 8-azabicyclo[3.2.1]octane.
112. The compound of claim 111 which is (2R)-1-(3-benzo[*d*]isoxazol-3-yl-8-azabicyclo[3.2.1]oct-8-yl-3-(3-thieno[2,3-*d*]isoxazol-3-yl-phenoxy)propan-2-ol.
113. A compound according to claim 1, wherein R_3 is adamantyl.
114. The compound of claim 113 which is (2R)-1-(adamantan-1-ylamino)-3-(3-thieno[2,3-*d*]isoxazol-3-yl-phenoxy)-propan-2-ol.
115. A compound according to claim 1, wherein Q is cyclohexyl.
116. compound of claim 115 which is (2R)-1-(cyclohexylmethyl-amino)-3-(3-thieno[2,3-*d*]isoxazol-3-yl-phenoxy)-propan-2-ol.
117. A compound according to claim 1, wherein Q is benzimidazolyl.

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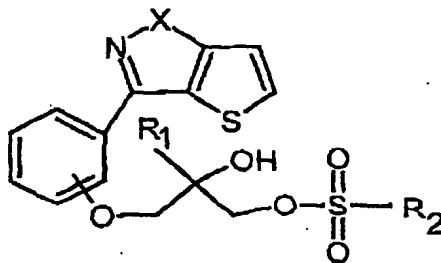
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118. The compound of claim 117 which is (2R)-1-[(1H-benzimidazol-2-ylmethyl)amino]-3-[3-thieno[2,3-d]isoxazol-3-yl-phenoxy]propan-2-ol.
119. A compound according to claim 1, wherein R_3 is 1,2,3,4-tetrahydronaphthyl.
120. The compound of claim 119 which is (2R)-1-(1,2,3,4-tetrahydronaphthalen-1-ylamino)-3-[3-thieno[2,3-d]isoxazol-3-yl]phenoxy]propan-2-ol.
121. A compound of formula



wherein X is N(CH₃) or O; and
 R_1 is C₁₋₆alkyl.

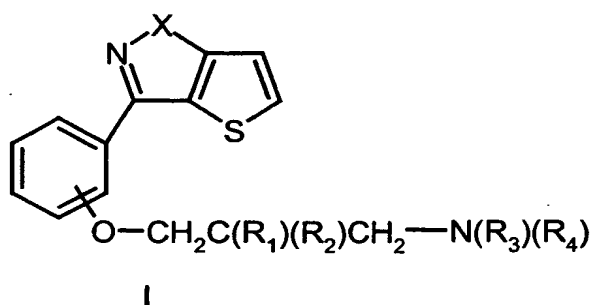
122. A compound of formula



wherein X is N(CH₃) or O;
 R_1 is H or C₁₋₆alkyl; and
 R_2 is CH₃, F, p-bromobenzene, p-nitrobenzene, or p-methylbenzene.

123. A method for antagonizing the effects of dopamine at the D₄ receptor comprising administering a compound according to claim 1 to a patient in need thereof.
124. A composition comprising a compound according to claim 1 in admixture with an inert carrier.
125. The composition according to claim 124, wherein said inert carrier is a pharmaceutical carrier.
126. A method of treating psychoses comprising administering to a patient in need thereof a therapeutically effective amount of a compound according to claim 1.

127. A method of treating Attention Deficit Hyperactivity Disorder comprising administering to a patient in need thereof a therapeutically effective amount of a compound according to claim 1.
- 5 128. A method of treating Obsessive-Compulsive Disorder comprising administering to a patient in need thereof a therapeutically effective amount of a compound according to claim 1.
129. A method of treating Substance Abuse comprising administering to a patient in need thereof a therapeutically effective amount of a compound according to claim 1.
- 10 130. A method of treating Substance Dependence comprising administering to a patient in need thereof a therapeutically effective amount of a compound according to claim 1.
131. A method of treating Parkinson's Disease comprising administering to a patient in need thereof a therapeutically effective amount of a compound according to claim 1.
- 15 132. A method of treating Parkinsonism comprising administering to a patient in need thereof a therapeutically effective amount of a compound according to claim 1.
133. A method of treating Tardive Dyskinesia comprising administering to a patient in need thereof a therapeutically effective amount of a compound according to claim 1.
- 20 134. A method of treating Gilles de la Tourette Syndrome comprising administering to a patient in need thereof a therapeutically effective amount of a compound according to claim 1.
- 25 135. A method of treating Conduct Disorder comprising administering to a patient in need thereof a therapeutically effective amount of a compound according to claim 1.
136. A method of treating Oppositional Defiant Disorder comprising administering to a patient in need thereof a therapeutically effective amount of a compound according to claim 1.
- 30 137. A method of making a compound of formula I



a pharmaceutically acceptable salt or stereoisomer thereof,

wherein

X is N(CH₃) or O;

R₁ is OH or C₁₋₆ alkoxy;

R₂ is H or C₁₋₆ alkyl;

R₃ is (CH₂)_n Q, CH₂CH(OH)Q, CH(CH₃)Q, 1,2,3,4-tetrahydronaphthyl, indanyl, or adamantyl, wherein

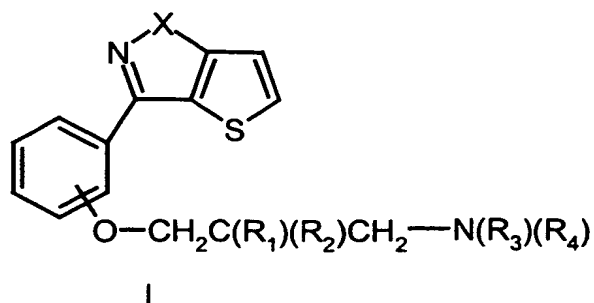
Q is thienyl, phenyl, furanyl, naphthyl, pyridyl, indolyl, indazolyl, cyclohexyl, 1,2-methylenedioxyphenyl, cyclohexenyl, 1H-pyrazolo[4,3-c]pyridyl, and

Q is optionally substituted with one or two moieties independently selected from halo, C₁₋₆ alkyl, C₁₋₆ alkoxy, hydroxy, S(O)₂NH₂, trifluoromethyl, or cyano, and

n is 1 or 2;

R₄ is H or C₁₋₆ alkyl; or

R₃ and R₄, together with the nitrogen atom to which R₃ and R₄ are attached, form 1,4-dioxo-8-azo-spiro[4.5]decanyl, piperazinyl, morpholinyl, piperidinyl, pyrrolidinyl, azocanyl, 1,2,3,4-tetrahydroisoquinoliny, 1,2,3,4-tetrahydro-β-carboliny, 4,5,6,7-tetrahydrothienyl[3,2-c]pyridyl, or 8-aza-bicyclo[3.2.1]octane, each of which may be mono- or independently di-substituted with halo, C₁₋₆ alkyl, C₁₋₆ alkoxy, C(O)phenyl, OH, CN, O-phenyl or (CH₂)_mZ,



a pharmaceutically acceptable salt or stereoisomer thereof,
wherein

X is N(CH₃) or O;

R₁ is OH or C₁₋₆ alkoxy;

R₂ is H or C₁₋₆ alkyl;

R₃ is (CH₂)_n Q, CH₂CH(OH)Q, CH(CH₃)Q, 1,2,3,4-tetrahydronaphthyl, indanyl, or adamantyl, wherein

Q is thienyl, phenyl, furanyl, naphthyl, pyridyl, indolyl, indazolyl,
cyclohexyl, 1,2-methylenedioxyphenyl, cyclohexenyl, 1H-
pyrazolo[4,3-c]pyridyl, and

Q is optionally substituted with one or two moieties independently
selected from halo, C₁₋₆ alkyl, C₁₋₆ alkoxy, hydroxy, S(O)₂NH₂,
trifluoromethyl, or cyano, and

n is 1 or 2;

R₄ is H or C₁₋₆ alkyl; or

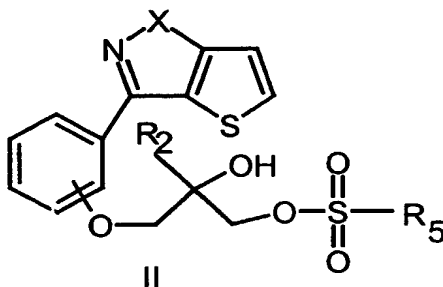
R₃ and R₄, together with the nitrogen atom to which R₃ and R₄ are attached, form
1,4-dioxo-8-azo-spiro[4.5]decanyl, piperazinyl, morpholinyl, piperidinyl,
pyrrolidinyl, azocanyl, 1,2,3,4-tetrahydroisoquinolinyl, 1,2,3,4-tetrahydro-β-
carbolinyl, 4,5,6,7-tetrahydrothienyl[3,2-c]pyridyl, or 8-aza-
bicyclo[3.2.1]octane, each of which may be mono- or independently di-
substituted with halo, C₁₋₆ alkyl, C₁₋₆ alkoxy, C(O)phenyl, OH, CN, O-phenyl or
(CH₂)_mZ,

Z is benzisoxazolyl, indazolyl, benzisothiazolyl, benzothienyl, pyrimidinyl, pyridyl, 1,2-methylenedioxyphenyl, or phenyl, and

Z, CH(OH)phenyl or O-phenyl are optionally substituted with one or two moieties independently selected from halo, C₁₋₆ alkyl, C₁₋₆ alkoxy, hydroxy, trifluoromethyl, S(O)₂NH₂, or cyano, and

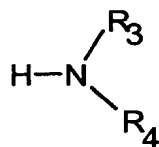
m is 0 or 1;

comprising the step of coupling a compound of formula II



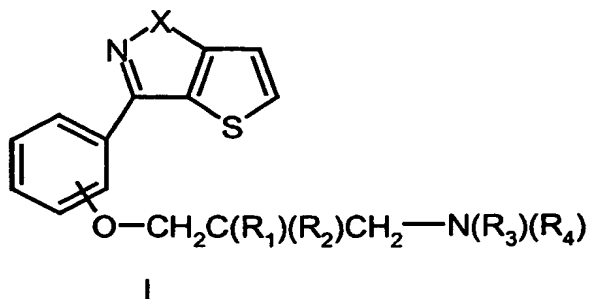
wherein X and R₂ are as defined in formula I; and
R₅ is CH₃, CF₃, F, p-bromobenzene, p-nitrobenzene, or p-methylbenzene;

with a reagent of formula III



wherein R₃ and R₄ are as defined in formula I;
to provide the compound of formula I.

139. A method of making a compound of formula I



a pharmaceutically acceptable salt or stereoisomer thereof,
wherein

X is N(CH₃) or O;

R₁ is OH or C₁₋₆ alkoxy;

5 R₂ is H or C₁₋₆ alkyl;

R₃ is (CH₂)_n Q, CH₂CH(OH)Q, CH(CH₃)Q, 1,2,3,4-tetrahydronaphthyl, indanyl, or
adamantyl, wherein

Q is thienyl, phenyl, furanyl, naphthyl, pyridyl, indolyl, indazolyl,
cyclohexyl, 1,2-methylenedioxyphenyl, cyclohexenyl, 1H-
10 pyrazolo[4,3-c]pyridyl, and

Q is optionally substituted with one or two moieties independently
selected from halo, C₁₋₆ alkyl, C₁₋₆ alkoxy, hydroxy, S(O)₂NH₂,
trifluoromethyl, or cyano, and

n is 1 or 2;

15

R₄ is H or C₁₋₆ alkyl; or

R₃ and R₄, together with the nitrogen atom to which R₃ and R₄ are attached, form
1,4-dioxo-8-azo-spiro[4.5]decanyl, piperaziny, morpholinyl, piperidinyl,
20 pyrrolidinyl, azocanyl, 1,2,3,4-tetrahydroisoquinolinyl, 1,2,3,4-tetrahydro-β-
carbolinyl, 4,5,6,7-tetrahydrothienyl[3,2-c]pyridyl, or 8-aza-
bicyclo[3.2.1]octane, each of which may be mono- or independently di-
substituted with halo, C₁₋₆ alkyl, C₁₋₆ alkoxy, C(O)phenyl, OH, CN, O-phenyl or
(CH₂)_mZ,

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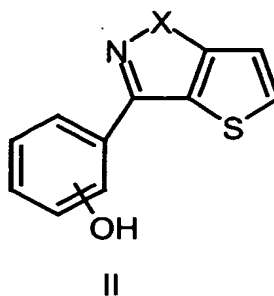
Z is benzisoxazolyl, indazolyl, benzisothiazolyl, benzothienyl,
pyrimidinyl, pyridyl, 1,2-methylenedioxyphenyl, or phenyl,
and

30

Z, CH(OH)phenyl or O-phenyl are optionally substituted with one
or two moieties independently selected from halo, C₁₋₆
alkyl, C₁₋₆ alkoxy, hydroxy, trifluoromethyl, S(O)₂NH₂, or
cyano, and

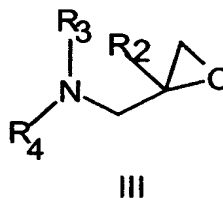
m is 0 or 1;

comprising the step of coupling a reagent of formula II



5 wherein X is as defined in formula I;

with a reagent of formula III



 wherein R₂, R₃, and R₄ are as defined in formula I;
to provide the compound of formula I.

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